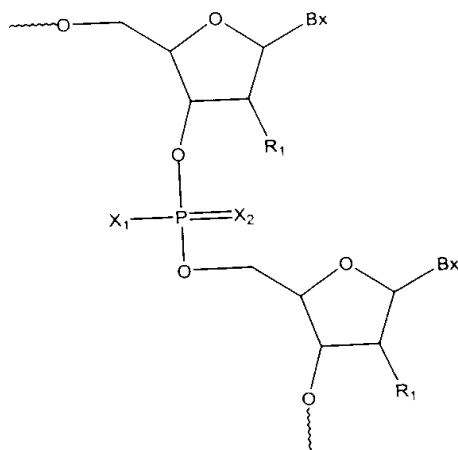


1. (Once Amended) A method of preparing an oligomeric compound having at least one moiety of formula:



wherein:

$X_2$  is O or S;

$X_1$  is Pg-O-, Pg-S-,  $C_1$ - $C_{10}$  straight or branched chain alkyl,  $CH_3(CH_2)_{nn}$ -O-,  $R_2R_3N$ - or a group remaining from coupling a chiral auxiliary;

$nn$  is from 0 to 10;

Pg is  $CH_3$ ,  $-CH_2CH_2CN$ ,  $-C(CH_3)(CH_3)CCl_3$ ,  $-CH_2CCl_3$ ,  $-CH_2CH=CH_2$ ,  $CH_2CH_2SiCH_3$ , 2-yl-ethyl phenylsulfonate,  $\delta$ -cyanobutenyl, cyano *p*-xylyl, diphenylsilylethyl, 4-nitro-2-yl-ethylbenzene, 2-yl-ethyl-methyl sulfonate, methyl-N-trifluoroacetyl ethyl, acetoxy phenoxy ethyl, or a blocking group;

$R_1$  is, independently, hydrogen, a blocked hydroxyl group, a sugar substituent group, a nitrogen protecting group, a substituted or unsubstituted  $C_1$ - $C_{10}$  alkyl, a substituted or unsubstituted  $C_2$ - $C_{10}$  alkenyl, or a substituted or unsubstituted  $C_2$ - $C_{10}$  alkynyl, wherein said

substitution is  $OR_3$ ,  $SR_3$ ,  $NH_3^+$ ,  $N(R_3)(R_4)$ , guanidine or acyl where said acyl is an acid amide or an ester;

$R_2$  is, independently, hydrogen, a  $C_1$ - $C_{10}$  alkyl, a cycloalkyl, an aryl, a nitrogen protecting group, a substituted or unsubstituted  $C_1$ - $C_{10}$  alkyl, a substituted or unsubstituted  $C_2$ - $C_{10}$  alkenyl, or a substituted or unsubstituted  $C_2$ - $C_{10}$  alkynyl, wherein said substitution is  $OR_3$ ,  $SR_3$ ,  $NH_3^+$ ,  $N(R_3)(R_4)$ , guanidine or acyl where said acyl is an acid amide or an ester;

or  $R_1$  and  $R_2$  together, are a nitrogen protecting group or are joined in a ring structure;

$R_3$  is, independently, hydrogen, a  $C_1$ - $C_{10}$  alkyl, a cycloalkyl, an aryl, or a nitrogen protecting group;

$R_4$  is, independently,  $N(L_1)L_2$ , hydrogen, a  $C_1$ - $C_{10}$  alkyl, or a nitrogen protecting group;

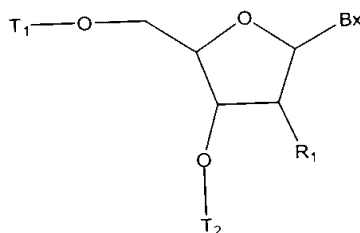
or  $R_3$  and  $R_4$ , together, are a nitrogen protecting group;

or  $R_3$  and  $R_4$  are joined in a ring structure;

or optionally,  $R_2$  and  $R_3$ , together with the nitrogen atom to which they are attached form a cyclic moiety;

each  $Bx$  is, independently, a heterocyclic base moiety; and  
comprising the steps of:

(a) providing a 5'-O-protected compound of the formula:



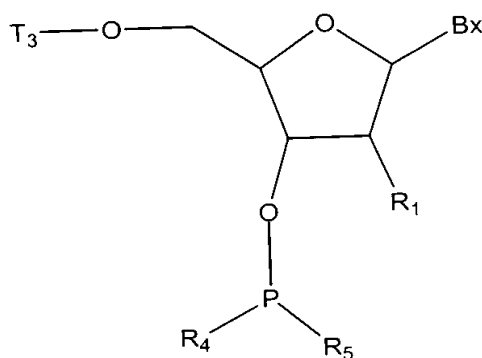
wherein:

T<sub>1</sub> is a hydroxyl protecting group; and

T<sub>2</sub> is a covalent attachment to a support media, a nucleoside bound to a support media, a nucleotide, an oligonucleoside or an oligonucleotide;

(b) treating said 5'-O-protected compound with a deprotecting reagent for a time and under conditions effective to form a 5'-O-deprotected compound;

(c) coupling said 5'-O-deprotected compound with an activated phosphorus composition of the formula:



wherein:

T<sub>3</sub> is a hydroxyl protecting group, a nucleoside, a nucleotide, an oligonucleoside or an oligonucleotide;

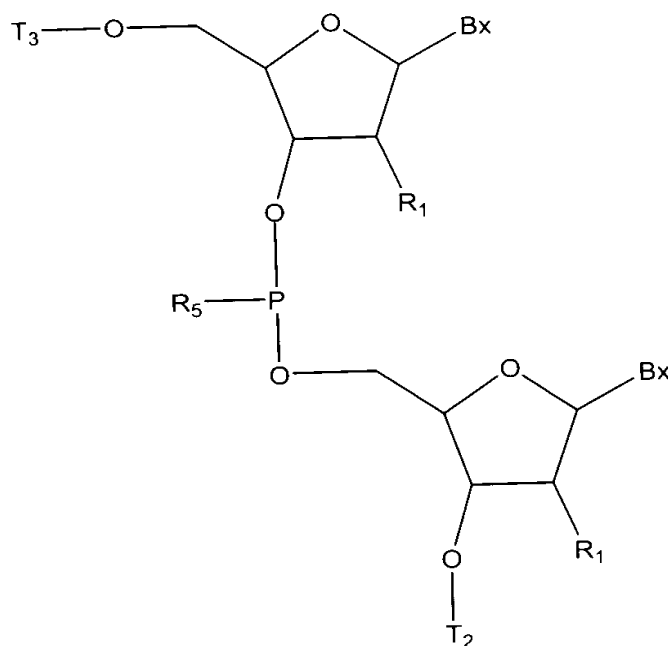
each L<sub>1</sub> and L<sub>2</sub> is, independently, C<sub>1-6</sub> straight or branched alkyl, or a C<sub>5-7</sub> cyclic aliphatic ring system;

or L<sub>1</sub> and L<sub>2</sub> are joined together to form a 4- to 13-membered heterocyclic ring system including the nitrogen atom to which L<sub>1</sub> and L<sub>2</sub> are attached; and

R<sub>5</sub> is X<sub>1</sub>;

or  $R_4$  and  $R_5$  together with the phosphorus atom to which  $R_4$  and  $R_5$  are attached form a chiral auxiliary;

for a time and under conditions effective to form an extended compound having the formula:

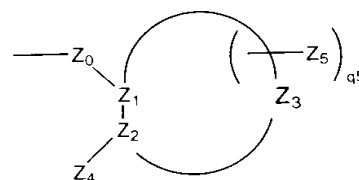


(d) treating said extended compound with a mixture comprising an oxidizing reagent and a capping reagent in a single step and for a time and under conditions effective to form said oligomeric compound.

27. (Once Amended) The method of claim 1 wherein each of said sugar substituent groups is, independently,  $C_1$ - $C_{20}$  alkyl,  $C_2$ - $C_{20}$  alkenyl,  $C_2$ - $C_{20}$  alkynyl,  $C_5$ - $C_{20}$  aryl, O-alkyl, O-alkenyl, O-alkynyl, O-aryl, O-aralkyl, O-alkylamino, O-alkylaminoalkyl (O-alkyl-N(H)alkyl), O-alkylaminodialkyl (O-alkyl-N-(alkyl)<sub>2</sub>), O-alkylalkoxy (O-alkyl-O-alkyl), O-alkyl-(N-imidazole), thiol, S-alkyl, S-alkenyl, S-alkynyl, NH-alkyl, NH-alkenyl, NH-alkynyl, N-dialkyl, S-aryl, NH-aryl, S-aralkyl, NH-aralkyl, N-phthalimido, halogen keto, carboxyl, nitro, nitroso,

$$Z_0 - \left\{ (CH_2)_{q_1} - O - \left( \overset{\overset{R_1}{|}}{N} \right)_{q_2} \right\}_{q_3} - (CH_2)_{q_4} - J - E$$

I

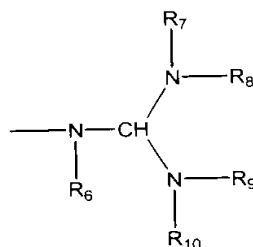


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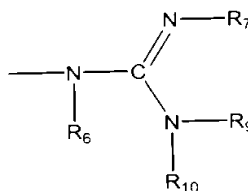
$Z_0$  is O, S or NH;

J is a single bond, O or C(=O);

E is C<sub>1</sub>-C<sub>10</sub> alkyl, N(R<sub>1</sub>)(R<sub>2</sub>), N(R<sub>1</sub>)(R<sub>5</sub>), N=C(R<sub>1</sub>)(R<sub>2</sub>), N=C(R<sub>1</sub>)(R<sub>5</sub>) or has one of formula III or IV;



III



IV

each R<sub>6</sub>, R<sub>7</sub>, R<sub>8</sub>, R<sub>9</sub> and R<sub>10</sub> is, independently, hydrogen, C(O)R<sub>11</sub>, substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, substituted or unsubstituted C<sub>2</sub>-C<sub>10</sub> alkenyl, substituted or unsubstituted C<sub>2</sub>-C<sub>10</sub> alkynyl, alkylsulfonyl, arylsulfonyl, a chemical functional group or a

conjugate group, wherein the substituent groups are selected from hydroxyl, amino, alkoxy, carboxy, benzyl, phenyl, nitro, thiol, thioalkoxy, halogen, alkyl, aryl, alkenyl and alkynyl;

or optionally, R<sub>7</sub> and R<sub>8</sub>, together form a phthalimido moiety with the nitrogen atom to which they are attached;

or optionally, R<sub>9</sub> and R<sub>10</sub>, together form a phthalimido moiety with the nitrogen atom to which they are attached;

each R<sub>11</sub> is, independently, substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, trifluoromethyl, cyanoethyloxy, methoxy, ethoxy, t-butoxy, allyloxy, 9-fluorenylmethoxy, 2-(trimethylsilyl)-ethoxy, 2,2,2-trichloroethoxy, benzyloxy, butyryl, iso-butyryl, phenyl or aryl;

R<sub>5</sub> is T-L,

T is a bond or a linking moiety;

L is a chemical functional group, a conjugate group or a support media;

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each R<sub>1</sub> and R<sub>2</sub> is, independently, H, a nitrogen protecting group, substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, substituted or unsubstituted C<sub>2</sub>-C<sub>10</sub> alkenyl, substituted or unsubstituted C<sub>2</sub>-C<sub>10</sub> alkynyl, wherein said substitution is OR<sub>3</sub>, SR<sub>3</sub>, NH<sub>3</sub><sup>+</sup>, N(R<sub>3</sub>)(R<sub>4</sub>), guanidino or acyl where said acyl is an acid amide or an ester;

or R<sub>1</sub> and R<sub>2</sub>, together, are a nitrogen protecting group or are joined in a ring structure that optionally includes an additional heteroatom selected from N and O;

or R<sub>1</sub>, T and L, together, are a chemical functional group;

each R<sub>3</sub> and R<sub>4</sub> is, independently, H, C<sub>1</sub>-C<sub>10</sub> alkyl, a nitrogen protecting group, or R<sub>3</sub> and R<sub>4</sub>, together, are a nitrogen protecting group;

or R<sub>3</sub> and R<sub>4</sub> are joined in a ring structure that optionally includes an additional heteroatom selected from N and O;

$Z_4$  is OX, SX, or  $N(X)_2$ ;

each X is, independently, H,  $C_1$ - $C_8$  alkyl,  $C_1$ - $C_8$  haloalkyl,  $C(=NH)N(H)R_5$ ,

$C(=O)N(H)R_5$  or  $OC(=O)N(H)R_5$ ;

$R_5$  is H or  $C_1$ - $C_8$  alkyl;

$Z_1$ ,  $Z_2$  and  $Z_3$  comprise a ring system having from about 4 to about 7 carbon atoms or having from about 3 to about 6 carbon atoms and 1 or 2 hetero atoms wherein said hetero atoms are selected from oxygen, nitrogen and sulfur and wherein said ring system is aliphatic, unsaturated aliphatic, aromatic, or saturated or unsaturated heterocyclic;

$Z_5$  is alkyl or haloalkyl having 1 to about 10 carbon atoms, alkenyl having 2 to about 10 carbon atoms, alkynyl having 2 to about 10 carbon atoms, aryl having 6 to about 14 carbon atoms,  $N(R_1)(R_2)OR_1$ , halo,  $SR_1$  or CN;

each  $q_1$  is, independently, an integer from 1 to 10;

each  $q_2$  is, independently, 0 or 1;

$q_3$  is 0 or an integer from 1 to 10;

$q_4$  is an integer from 1 to 10;

$q_5$  is from 0, 1 or 2; and

provided that when  $q_3$  is 0,  $q_4$  is greater than 1.

35. (Once Amended) The method of claim 34 wherein said deprotecting reagent is dichloroacetic acid, trichloroacetic acid, zinc bromide,  $AlCl_3$ ,  $TiCl_4$ ,  $(Et)AlCl$ ,  $(i-Bu)_2AlCl$ , ceric ammonium nitrate, 1,1,1,3,3,3-hexafluoro-2-propanol or diethyloxomalonate.